Appl. No. 09/720,086; 102(e): July 23, 2001 Dkt. No. 0609.4560002/JAG/KRM/DJN; Group Art Unit: 1642 Inventors: Li et al.; Tel: 202/371-2600 Täle: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

### Mouse Dnmt3a DNA sequence



1	GAATTCCGGC CTGCTGCCGC	GCCGCCCGAC	CCGCCGGGCC	ACACGGCAGA
51	GCCGCCTGAA GCCCAGCGC	T GAGGCTGCAC	TTTTCCGAGG	GCTTGACATC
101	AGGGTCTATG TTTAAGTCT	T AGCTCTTGCT	TACAAAGACC	ACGCCAATTC
151	CTTCTCTGAA GCCCTCGCAC	CCCCACAGCG	CCCTCGCAGC	CCCAGCCTGC
201	CGCCTACTGC CCAGCAATGO	CCTCCAGCGG	CCCCGGGGAC	ACCAGCAGCT
251	CCTCTCTGGA GCGGGAGGA	T GATCGAAAGG	AAGGAGAGGA	ACAGGAGGAG
301	AACCGTGGCA AGGAAGAGCC	CCAGGAGCCC	AGCGCCACGG	CCCGGAAGGT
351	GGGGAGGCCT GGCCGGAAGG	GCAAGCACCC	ACCGGTGGAA	AGCAGTGACA
401	CCCCCAAGGA CCCAGCAGTO	ACCACCAAGT	CTCAGCCCAT	GGCCCAGGAC
451	TCTGGCCCCT CAGATCTGC	T ACCCAATGGA	GACTTGGAGA	AGCGGAGTGA
501	ACCCCAACCT GAGGAGGGG	A GCCCAGCTGC	AGGGCAGAAG	GGTGGGGCCC
551	CAGCTGAAGG AGAGGGAAC	GAGACCCCAC	CAGAAGCCTC	CAGAGCTGTG
601	GAGAATGGCT GCTGTGTGAG	CAAGGAAGGC	CGTGGAGCCT	CTGCAGGAGA
651	GGGCAAAGAA CAGAAGCAGA	A CCAACATCGA	ATCCATGAAA	ATGGAGGGCT
701	CCCGGGGCCG ACTGCGAGG	T GGCTTGGGCT	GGGAGTCCAG	CCTCCGTCAG
751	CGACCCATGC CAAGACTCAG	CTTCCAGGCA	GGGGACCCCT	ACTACATCAG
801	CAAACGGAAA CGGGATGAG	F GGCTGGCACG	TTGGAAAAGG	GAGGCTGAGA
851	AGAAAGCCAA GGTAATTGCA	A GTAATGAATG	CTGTGGAAGA	GAACCAGGCC
901	TCTGGAGAGT CTCAGAAGG	r ggaggaggcc	AGCCCTCCTG	CTGTGCAGCA
951	GCCCACGGAC CCTGCTTCTC	CGACTGTGGC	CACCACCCCT	GAGCCAGTAG
1001	GAGGGGATGC TGGGGACAAG	S AATGCTACCA	AAGCAGCCGA	CGATGAGCCT
1051	GAGTATGAGG ATGGCCGGGG	CTTTGGCATT	GGAGAGCTGG	TGTGGGGGAA
1101	ACTICGCGGC TICTCCTGG	T GGCCAGGCCG	AATTGTGTCT	TGGTGGATGA

ACL ASS

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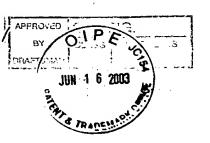
CAGGCCGGAG CCGAGCAGCT GAAGGCACTC GCTGGGTCAT GTGGTTCGGA 1151 GATGGCAAGT TCTCAGTGGT GTGTGTGGAG AAGCTCATGC CGCTGAGCTC 1201 CTTCTGCAGT GCATTCCACC AGGCCACCTA CAACAAGCAG CCCATGTACC GCAAAGCCAT CTACGAAGTC CTCCAGGTGG CCAGCAGCCG TGCCGGGAAG 1301 CTGTTTCCAG CTTGCCATGA CAGTGATGAA AGTGACAGTG GCAAGGCTGT GGAAGTGCAG AACAAGCAGA TGATTGAATG GGCCCTCGGT GGCTTCCAGC CCTCGGGTCC TAAGGGCCTG GAGCCACCAG AAGAAGAGAA GAATCCTTAC AAGGAAGTTT ACACCGACAT GTGGGTGGAG CCTGAAGCAG CTGCTTACGC 1501 CCCACCCCA CCAGCCAAGA AACCCAGAAA GAGCACAACA GAGAAACCTA AGGTCAAGGA GATCATTGAT GAGCGCACAA GGGAGCGGCT GGTGTATGAG 1601 GTGCGCCAGA AGTGCAGAAA CATCGAGGAC ATTTGTATCT CATGTGGGAG CCTCAATGTC ACCCTGGAGC ACCCACTCTT CATTGGAGGC ATGTGCCAGA 1701 ACTGTAAGAA CTGCTTCTTG GAGTGTGCTT ACCAGTATGA CGACGATGGG TACCAGTCCT ATTGCACCAT CTGCTGTGGG GGGCGTGAAG TGCTCATGTG TGGGAACAAC AACTGCTGCA GGTGCTTTTG TGTCGAGTGT GTGGATCTCT 1851 TGGTGGGGCC AGGAGCTGCT CAGGCAGCCA TTAAGGAAGA CCCCTGGAAC 1901 TGCTACATGT GCGGGCATAA GGGCACCTAT GGGCTGCTGC GAAGACGGGA 1951 AGACTGGCCT TCTCGACTCC AGATGTTCTT TGCCAATAAC CATGACCAGG 2001 AATTTGACCC CCCAAAGGTT TACCCACCTG TGCCAGCTGA GAAGAGGAAG 2051 CCCATCCGCG TGCTGTCTCT CTTTGATGGG ATTGCTACAG GGCTCCTGGT GCTGAAGGAC CTGGGCATCC AAGTGGACCG CTACATTGCC TCCGAGGTGT GTGAGGACTC CATCACGGTG GGCATGGTGC GGCACCAGGG AAAGATCATG TACGTCGGG ACGTCCGCAG CGTCACACAG AAGCATATCC AGGAGTGGGG 2251 CCCATTCGAC CTGGTGATTG GAGGCAGTCC CTGCAATGAC CTCTCCATTG



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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

. 2351 TCAACCCTGC CCGCAAGGGA CITTATGAGG GTACTGGCCG CCTCTTC

2351	TCAACCCTGC	CCGCAAGGGA	CTTTATGAGG	GTACTGGCCG	CCTCTTCTTT
2401	GAGTTCTACC	GCCTCCTGCA	TGATGCGCGG	CCCAAGGAGG	GAGATGATCG
2451	CCCCTTCTTC	TGGCTCTTTG	AGAATGTGGT	GGCCATGGGC	GTTAGTGACA
2501	AGAGGGACAT	CTCGCGATTT	CTTGAGTCTA	ACCCCGTGAT	GATTGACGCC
2551	AAAGAAGTGT	CTGCTGCACA	CAGGGCCCGT	TACTTCTGGG	GTAACCTTCC
2601	TGGCATGAAC	AGGCCTTTGG	CATCCACTGT	GAATGATAAG	CTGGAGCTGC
2651	AAGAGTGTCT	GGAGCACGGC	AGAATAGCCA	AGTTCAGCAA	AGTGAGGACC
2701	ATTACCACCA	GGTCAAACTC	TATAAAGCAG	GGCAAAGACC	AGCATTTCCC
2751	CGTCTTCATG	AACGAGAAGG	AGGACATCCT	GTGGTGCACT	GAAATGGAAA
2801	GGGTGTTTGG	CTTCCCCGTC	CACTACACAG	ACGTCTCCAA	CATGAGCCGC
2851	TTGGCGAGGC	AGAGACTGCT	GGGCCGATCG	TGGAGCGTGC	CGGTCATCCG
2901	CCACCTCTTC	GCTCCGCTGA	AGGAATATTT	TGCTTGTGTG	TAAGGGACAT
2951	GGGGCAAAC	TGAAGTAGTG	ATGATAAAAA	AGTTAAACAA	ACAAACAAAC
3001	AAAAAACAAA	ACAAAACAAT	AAAACACCAA	GAACGAGAGG	ACGGAGAAAA
3051	GTTCAGCACC	CAGAAGAGAA	AAAGGAATTT	AAAGCAAACC	ACAGAGGAGG
3101	AAAACGCCGG	AGGGCTTGGC	CTTGCAAAAG	GGTTGGACAT	CATCTCCTGA
3151	GTTTTCAATG	TTAACCTTCA	GTCCTATCTA	AAAAGCAAAA	TAGGCCCCTC
3201	CCCTTCTTCC	CCTCCGGTCC	TAGGAGGCGA	ACTITITGTT	TTCTACTCTT
3251	TTTCAGAGGG	GTTTTCTGTT	TGTTTGGGTT	TTTGTTTCTT	GCTGTGACTG
3301	AAACAAGAGA	GTTATTGCAG	CAAAATCAGT	AACAACAAAA	AGTAGAAATG
3351	CCTTGGAGAG	GAAAGGGAGA	GAGGGAAAAT	TCTATAAAAA	CTTAAAATAT
3401	TGGTTTTTT	TTTTTTCCT	TTTCTATATA	TCTCTTTGGT	TGTCTCTAGC
3451	CTGATCAGAT	AGGAGCACAA	ACAGGAAGAG	AATAGAGACC	CTCGGAGGCA
3501	GAGTCTCCTC	TCCCACCCCC	CGAGCAGTCT	CAACAGCACC	ATTCCTGGTC



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Title: De Novo DNA Cytosine Methyltransferase Genea, Polypeptides
and Meer Thereof

3551 AGACACTITC TACAGTATTT CAGGTGCCTA CCACACAGGA AACCTTGAAG 3601 AAAACCAGTT TCTAGAAGCC GCTGTTACCT CTTGTTTACA GTTTATATAT 3651 ATATGATAGA TATGAGATAT ATATATAA AAGGTACTGT TAACTACTGT 3701 ACATCCCGAC TTCATAATGG TGCTTTCAAA ACAGCGAGAT GAGCAAAGAC 3751 3801 ATCAGCTTCC GCCTGGCCCT CTGTGCAAAG GGTTTCAGCC CAGGATGGGG 3851 AGAGGGGAGC AGCTGGAGGG GGTTTTAACA AACTGAAGGA TGACCCATAT 3901 CACCCCCAC CCCTGCCCCA TGCCTAGCTT CACCTGCCAA AAAGGGGCTC 3951 AGCTGAGGTG GTCGGACCCT GGGGAAGCTG AGTGTGGAAT TTATCCAGAC TCGCGTGCAA TAACCTTAGA ATATGAATCT AAAATGACTG CCTCAGAAAA 4001 ATGGCTTGAG AAAACATTGT CCCTGATTTT GAATTCGTCA GCCACGTTGA 4051 AGGCCCCTTG TGGGATCAGA AATATTCCAG AGTGAGGGAA AGTGACCCGC 4101 CATTAACCCC NCCTGGAGCA AATAAAAAA CATACAAAAT GT



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Title: De Novo DNA Cytosine Methyltransferase Genea, Polypeptides
and Uses Thereof

#### Mouse Dnmt3b1 DNA Sequence

GAATTCCGGG CGCCGGGGTT AAGCGGCCCA AGTAAACGTA GCGCAGCGAT CGGCGCCGGA GATTCGCGAA CCCGACACTC CGCGCCGCCC GCCGGCCAGG 101 ACCCGCGGC CGATCGCGC GCCGCGCTAC AGCCAGCCTC ACGACAGGCC CGCTGAGGCT TGTGCCAGAC CTTGGAAACC TCAGGTATAT ACCTTTCCAG 151 ACGCGGGATC TCCCCTCCCC CATCCATAGT GCCTTGGGAC CAAATCCAGG 201 GCCTTCTTC AGGAAACAAT GAAGGGAGAC AGCAGACATC TGAATGAAGA 251 AGAGGGTGCC AGCGGGTATG AGGAGTGCAT TATCGTTAAT GGGAACTTCA 301 351 GTGACCAGTC CTCAGACACG AAGGATGCTC CCTCACCCCC AGTCTTGGAG 401 GCAATCTGCA CAGAGCCAGT CTGCACACCA GAGACCAGAG GCCGCAGGTC 451 AAGCTCCCGG CTGTCTAAGA GGGAGGTCTC CAGCCTTCTG AATTACACGC AGGACATGAC AGGAGATGGA GACAGAGATG ATGAAGTAGA TGATGGGAAT 501 551 GGCTCTGATA TTCTAATGCC AAAGCTCACC CGTGAGACCA AGGACACCAG GACGCGCTCT GAAAGCCCGG CTGTCCGAAC CCGACATAGC AATGGGACCT 601 CCAGCTTGGA GAGGCAAAGA GCCTCCCCCA GAATCACCCG AGGTCGGCAG 651 GGCCGCCACC ATGTGCAGGA GTACCCTGTG GAGTTTCCGG CTACCAGGTC 701 751 TCGGAGACGT CGAGCATCGT CTTCAGCAAG CACGCCATGG TCATCCCCTG CCAGCGTCGA CTTCATGGAA GAAGTGACAC CTAAGAGCGT CAGTACCCCA 801 TCAGTTGACT TGAGCCAGGA TGGAGATCAG GAGGGTATGG ATACCACACA 851 901 GGTGGATGCA GAGAGCAGAG ATGGAGACAG CACAGAGTAT CAGGATGATA AAGAGTTTGG AATAGGTGAC CTCGTGTGGG GAAAGATCAA GGGCTTCTCC 951 TGGTGGCCTG CCATGGTGGT GTCCTGGAAA GCCACCTCCA AGCGACAGGC 1001

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CATGCCCGGA ATGCGCTGGG TACAGTGGTT TGGTGATGGC AAGTTTTCTG 1051 AGATCTCTGC TGACAAACTG GTGGCTCTGG GGCTGTTCAG CCAGCACTTT 1101 AATCTGGCTA CCTTCAATAA GCTGGTTTCT TATAGGAAGG CCATGTACCA 1151 CACTCTGGAG AAAGCCAGGG TTCGAGCTGG CAAGACCTTC TCCAGCAGTC 1201 1251 CTGGAGAGTC ACTGGAGGAC CAGCTGAAGC CCATGCTGGA GTGGGCCCAC GGTGGCTTCA AGCCTACTGG GATCGAGGGC CTCAAACCCA ACAAGAAGCA 1301 ACCAGTGGTT AATAAGTCGA AGGTGCGTCG TTCAGACAGT AGGAACTTAG 1351 AACCCAGGAG ACGCGAGAAC AAAAGTCGAA GACGCACAAC CAATGACTCT GCTGCTTCTG AGTCCCCCCC ACCCAAGCGC CTCAAGACAA ATAGCTATGG 1501 CGGGAAGGAC CGAGGGGAGG ATGAGGAGAG CCGAGAACGG ATGGCTTCTG 1551 AAGTCACCAA CAACAAGGGC AATCTGGAAG ACCGCTGTTT GTCCTGTGGA AAGAAGAACC CTGTGTCCTT CCACCCCCTC TTTGAGGGTG GGCTCTGTCA GAGTTGCCGG GATCGCTTCC TAGAGCTCTT CTACATGTAT GATGAGGACG GCTATCAGTC CTACTGCACC GTGTGCTGTG AGGGCCGTGA ACTGCTGCTG 1701 TGCAGTAACA CAAGCTGCTG CAGATGCTTC TGTGTGGAGT GTCTGGAGGT 1751 GCTGGTGGGC GCAGGCACAG CTGAGGATGC CAAGCTGCAG GAACCCTGGA GCTGCTATAT GTGCCTCCCT CAGCGCTGCC ATGGGGTCCT CCGACGCAGG AAAGATTGGA ACATGCGCCT GCAAGACTTC TTCACTACTG ATCCTGACCT 1901 GGAAGAATTT GAGCCACCCA AGTTGTACCC AGCAATTCCT GCAGCCAAAA GGAGGCCCAT TAGAGTCCTG TCTCTGTTTG ATGGAATTGC AACGGGGTAC 2001 TIGGIGCICA AGGACTIGGG TATTAAAGIG GAAAAGIACA TIGCCICCGA 2051 AGTCTGTGCA GAGTCCATCG CTGTGGGAAC TGTTAAGCAT GAAGGCCAGA 2101 TCAAATATGT CAATGACGTC CGGAAAATCA CCAAGAAAAA TATTGAAGAG 2151 TGGGGCCCGT TCGACTTGGT GATTGGTGGA AGCCCATGCA ATGATCTCTC 2201

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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

2251	TAACGTCAAT	CCTGCCCGCA	AAGGTTTATA	TGAGGGCACA	GGAAGGCTCT
2301	TCTTCGAGTT	TTACCACTTG	CTGAATTATA	CCCGCCCCAA	GGAGGGCGAC
2351	AACCGTCCAT	TCTTCTGGAT	GTTCGAGAAT	GTTGTGGCCA	TGAAAGTGAA
2401	TGACAAGAAA	GACATCTCAA	GATTCCTGGC	ATGTAACCCA	GTGATGATCG
2451	ATGCCATCAA	GGTGTCTGCT	GCTCACAGGG	CCCGGTACTT	CTGGGGTAAC
2501	CTACCCGGAA	TGAACAGGCC	CGTGATGGCT	TCAAAGAATG	ATAAGCTCGA
2551	GCTGCAGGAC	TGCCTGGAGT	TCAGTAGGAC	AGCAAAGTTA	AAGAAAGTGC
2601	AGACAATAAC	CACCAAGTCG	AACTCCATCA	GACAGGGCAA	AAACCAGCTT
2651	TTCCCTGTAG	TCATGAATGG	CAAGGACGAC	GTTTTGTGGT	GCACTGAGCT
2701	CGAAAGGATC	TTCGGCTTCC	CTGCTCACTA	CACGGACGTG	TCCAACATGG
2751	GCCGCGGCGC	CCGTCAGAAG	CTGCTGGGCA	GGTCCTGGAG	TGTACCGGTC
2801	ATCAGACACC	TGTTTGCCCC	CTTGAAGGAC	TACTITICCCT	GTGAATAGTT
2851	CTACCCAGGA	CTGGGGAGCT	CTCGGTCAGA	GCCAGTGCCC	AGAGTCACCC
2901	CTCCCTGAAG	GCACCTCACC	TGTCCCCTTT	TTAGCTCACC	TGTGTGGGGC
2951	CTCACATCAC	TGTACCTCAG	CTTTCTCCTG	CTCAGTGGGA	GCAGAGCCTC
3001	CTGGCCCTTG	CAGGGGAGCC	CCGGTGCTCC	CTCCGTGTGC	ACAGCTCAGA
3051	CCTGGCTGCT	TAGAGTAGCC	CGGCATGGTG	CTCATGTTCT	CTTACCCTGA
3101	AACTTTAAAA	CTTGAAGTAG	GTAGTAAGAT	GGCTTTCTTT	TACCCTCCTG
3151	AGTTTATCAC	TCAGAAGTGA	TGGCTAAGAT	ACCAAAAAA	CAAACAAAAA
3201	CAGAAACAAA	AAACAAAAA	AAACCTCAAC	AGCTCTCTTA	GTACTCAGGT
3251	TCATGCTGCA	AAATCACTTG	AGATTTTGTT	TTTAAGTAAC	CCGTGCTCCA
3301	CATTIGCTGG	AGGATGCTAT	TGTGAATGTG	GGCTCAGATG	AGCAAGGTCA
3351	AGGGGCCAAA	AAAAATTCCC	CCTCTCCCCC	CAGGAGTATT	TGAAGATGAT
3401	GTTTATGGTT	TAAGTCTTCC	TGGCACCTTC	CCCTTGCTTT	GGTACAAGGG



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Title: De Novo DNA Cytosine Methyttransferase Genes, Polypeptide

CTGAAGTCCT GTTGGTCTTG TAGCATTTCC CAGGATGATG ATGTCAGCAG 3501 GGATGACATC ACCACCTTTA GGGCTTTTCC CTGGCAGGGG CCCATGTGGC TAGTCCTCAC GAAGACTGGA GTAGAATGTT TGGAGCTCAG GAAGGGTGGG 3551 TGGAGTGGCC CTCTTCCAGG TGTGAGGGAT ACGAAGGAGG AAGCTTAGGG 3601 AAATCCATTC CCCACTCCCT CTTGCCAAAT GAGGGGCCCA GTCCCCAACA 3651 GCTCAGGTCC CCAGAACCCC CTAGTTCCTC ATGAGAAGCT AGGACCAGAA 3751 GCACATCGTT CCCCTTATCT GAGCAGTGTT TGGGGAACTA CAGTGAAAAC 3801 CTTCTGGAGA TGTTAAAAGC TTTTTACCCC ACGATAGATT GTGTTTTTAA 3851 GGGGTGCTTT TTTTAGGGGC ATCACTGGAG ATAAGAAAGC TGCATTTCAG AAATGCCATC GTAATGGTTT TTAAACACCT TTTACCTAAT TACAGGTGCT 3901 3951 ATTITATAGA AGCAGACAAC ACTICTITIT ATGACTCTCA GACTICTATT 4001 TICATGTTAC CATTITITIT GTAACTCGCA AGGTGTGGGC TITTGTAACT 4051 TCACAGGTGT GGGGAGAGAC TGCCTTGTTT CAACAGTTTG TCTCCACTGG TITCTAATTT TTAGGTGCAA AGATGACAGA TGCCCAGAGT TTACCTTTCT 4101 4151

FIG. 1B-4

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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides
and Liese: Thereof

#### Human DNMT3A DNA Sequence

GCCGCGG CACCAGGGCG CGCAGCCGGG 1 CCGCCCGAC CCCACCGCC ATACGGTGGA GCCATCGAAG CCCCCACCCA CAGGCTGACA GAGGCACCGT TCACCAGAGG GCTCAACACC GGGATCTATG TTTAAGTTTT AACTCTCGCC TCCAAAGACC ACGATAATTC CTTCCCCAAA 128 GCCCAGCAGC CCCCCAGCCC CGCGCAGCCC CAGCCTGCCT CCCGGCGCCCC 178 AGATGCCCGC CATGCCCTCC AGCGGCCCCG GGGACACCAG CAGCTCTGCT 228 GCGGAGCGGG AGGAGGACCG AAAGGACGGA GAGGAGCAGG AGGAGCCGCG 278 328 TGGCAAGGAG GAGCGCCAAG AGCCCAGCAC CACGGCACGG AAGGTGGGGC GGCCTGGGAG GAAGCGCAAG CACCCCCGG TGGAAAGCGG TGACACGCCA 378 AAGGACCCTG CGGTGATCTC CAAGTCCCCA TCCATGGCCC AGGACTCAGG 428 CGCCTCAGAG CTATTACCCA ATGGGGACTT GGAGAAGCGG AGTGAGCCCC 478 AGCCAGAGGA GGGGAGCCCT GCTGGGGGGC AGAAGGGCGG GGCCCCAGCA 528 578 GAGGGAGAGG GTGCAGCTGA GACCCTGCCT GAAGCCTCAA GAGCAGTGGA AAATGGCTGC TGCACCCCCA AGGAGGGCCG AGGAGCCCCT GCAGAAGCGG 628 GCAAAGAACA GAAGGAGACC AACATCGAAT CCATGAAAAT GGAGGGCTCC 678 CGGGGCCGGC TGCGGGGTGG CTTGGGCTGG GAGTCCAGCC TCCGTCAGCG 728 GCCCATGCCG AGGCTCACCT TCCAGGCGGG GGACCCCTAC TACATCAGCA AGCGCAAGCG GGACGAGTGG CTGGCACGCT GGAAAAGGGA GGCTGAGAAG 828 AAAGCCAAGG TCAGTGCAGG AATGAATGCT GTGGAAGAAA ACCAGGGGCC 878 CGGGGAGTCT CAGAAGGTGG AGGAGGCCAG CCCTCCTGCT GTGCAGCAGC 928 CCACTGACCC CGCATCCCCC ACTGTGGCTA CCACGCCTGA GCCCGTGGGG TCCGATGCTG GGGACAAGAA TGCCACCAAA GCAGGCGATG ACGAGCCAGA 1028



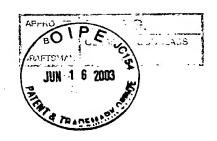
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GTACGAGGAC GGCCGGGGT TTGGCATTGG GGAGCTGGTG TGGGGGAAAC TGCGGGGCTT CTCCTGGTGG CCAGGCCGCA TTGTGTCTTG GTGGATGACG 1128 GGCCGGAGCC GAGCAGCTGA AGGCACCCGC TGGGTCATGT GGTTCGGAGA 1178 CGGCAAATTC TCAGTGGTGT GTGTTGAGAA GCTGATGCCG CTGAGCTCGT 1228 TITGCAGTGC GTTCCACCAG GCCACGTACA ACAAGCAGCC CATGTACCGC 1278 AAAGCCATCT ACGAGGTCCT GCAGGTGGCC AGCAGCCGCG CGGGGAAGCT 1328 GTTCCCGGTG TGCCACGACA GCGATGAGAG TGACACTGCC AAGGCCGTCG 1378 1428 AGGTGCAGAA CAAGCCCATG ATTGAATGGG CCCTGGGGGG CTTCCAGCCT 1478 AGAAGTGTAC ACGGACATGT GGGTGGAACC TGAGGCAGCT GCCTACGCAC 1528 CACCTCCACC AGCCAAAAAG CCCCGGAAGA GCACAGCGGA GAAGCCCAAG 1578 GTCAAGGAGA TTATTGATGA GCGCACAAGA GAGCGGCTGG TGTACGAGGT 1628 GCGCAGAAG TGCCGGAACA TTGAGGACAT CTGCATCTCC TGTGGGAGCC TCAATGTTAC CCTGGAACAC CCCCTCTTCG TTGGAGGAAT GTGCCAAAAC 1728 1778 TGCAAGAACT GCTTTCTGGA GTGTGCGTAC CAGTACGACG ACGACGGCTA CCAGTCCTAC TGCACCATCT GCTGTGGGGG CCGTGAGGTG CTCATGTGCG 1828 GAAACAACAA CTGCTGCAGG TGCTTTTGCG TGGAGTGTGT GGACCTCTTG GTGGGGCCGG GGGCTGCCCA GGCAGCCATT AAGGAAGACC CCTGGAACTG 1928 1978 CTACATGTGC GGGCACAAGG GTACCTACGG GCTGCTGCGG CGGCGAGAGG ACTGGCCCTC CCGGCTCCAG ATGTTCTTCG CTAATAACCA CGACCAGGAA 2028 2078 TITGACCCTC CAAAGGTTTA CCCACCTGTC CCAGCTGAGA AGAGGAAGCC 2128 CATCCGGGTG CTGTCTCTCT TTGATGGAAT CGCTACAGGG CTCCTGGTGC TGAAGGACTT GGGCATTCAG GTGGACCGCT ACATTGCCTC GGAGGTGTGT 2178

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GAGGACTCCA TCACGGTGGG CATGGTGCGG CACCAGGGGA AGATCATGTA 2228 CGTCGGGGAC GTCCGCAGCG TCACACAGAA GCATATCCAG GAGTGGGGCC CATTCGATCT GGTGATTGGG GGCAGTCCCT GCAATGACCT CTCCATCGTC AACCCTGCTC GCAAGGGCCT CTACGAGGGC ACTGGCCGGC TCTTCTTTGA 2378 GTTCTACCGC CTCCTGCATG ATGCGCGGCC CAAGGAGGGA GATGATCGCC CCTTCTTCTG GCTCTTTGAG AATGTGGTGG CCATGGGCGT TAGTGACAAG AGGGACATCT CGCGATTTCT CGAGTCCAAC CCTGTGATGA TTGATGCCAA 2528 AGAAGTGTCA GCTGCACACA GGGCCCGCTA CTTCTGGGGT AACCTTCCCG 2578 GTATGAACAG GCCGTTGGCA TCCACTGTGA ATGATAAGCT GGAGCTGCAG 2678 GAGTGTCTGG AGCATGGCAG GATAGCCAAG TTCAGCAAAG TGAGGACCAT TACTACGAGG TCAAACTCCA TAAAGCAGGG CAAAGACCAG CATTTTCCTG 2728 2778 TCTTCATGAA TGAGAAAGAG GACATCTTAT GGTGCACTGA AATGGAAAGG GTATTTGGTT TCCCAGTCCA CTATACTGAC GTCTCCAACA TGAGCCGCTT GGCGAGGCAG AGACTGCTGG GCCGGTCATG GAGCGTGCCA GTCATCCGCC 2878 ACCTCTTCGC TCCGCTGAAG GAGTATTTTG CGTGTGTGTA AGGGACATGG 2928 GGGCAAACTG AGGTAGCGAC ACAAAGTTAA ACAAACAAAC AAAAAAACACA 2978 AAACATAATA AAACACCAAG AACATGAGGA TGGAGAGAAG TATCAGCACC 3028 3078 CAGAAGAGAA AAAGGAATTT AAAACAAAAA CCACAGAGGC GGAAATACCG GAGGGCTTTG CCTTGCGAAA AGGGTTGGAC ATCATCTCCT GATTTTTCAA 3128 3178 TCCCCCTTCC CTTTTTTTC GGTCAGACCT TITATTTTCT ACTCTTTCA 3228 GAGGGGTTTT CTGTTTGTTT GGGTTTTGTT TCTTGCTGTG ACTGAAACAA 3278 GAAGGTTATT GCAGCAAAAA TCAGTAACAA AAAATAGTAA CAATACCTTG 3378 CAGAGGAAAG GTGGGAGGAG AGGAAAAAAG GGAAATTTTT AAAGAAATCT



Dkt. No. 0609.4560002/JAG/KRM/DJN; Group Art Unit: 1642 Inventors: Li et al.; Tel: 202/371-2600

Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides

ATATATIGGG TIGITITITT TITTGTTTTT TGTTTTTTT TTTTGGGTTT TITITITIA CTATATATCI TITITITGII GICTCTAGCC IGATCAGATA GGAGCACAAG CAGGGGACGG AAAGAGAGAG ACACTCAGGC GGCAGCATTC 3528 CCTCCCAGCC ACTGAGCTGT CGTGCCAGCA CCATTCCTGG TCACGCAAAA CAGAACCCAG TTAGCAGCAG GGAGACGAGA ACACCACACA AGACATTTTT CTACAGTATT TCAGGTGCCT ACCACACAGG AAACCTTGAA GAAAATCAGT TICTAGAAGC CGCTGTTACC TCTTGTTTAC AGTTTATATA TATATGATAG ATATGAGATA TATATAAA AGGTACTGTT AACTACTGTA CAACCCGACT TCATAATGGT GCTTTCAAAC AGCGAGATGA GTAAAAACAT CAGCTTCCAC 3828 GTTGCCTTCT GCGCAAAGGG TTTCACCAAG GATGGAGAAA GGGAGACAGC TTGCAGATGG CGCGTTCTCA CGGTGGGCTC TTCCCCTTGG TTTGTAACGA 3928 AGTGAAGGAG GAGAACTTGG GAGCCAGGTT CTCCCTGCCA AAAAGGGGGGC 3978 4028 TAGATGAGGT GGTCGGGCCC GTGGACAGCT GAGAGTGGGA ITCATCCAGA CTCATGCAAT AACCCTTTGA TTGTTTTCTA AAAGGAGACT CCCTCGGCAA GATGGCAGAG GGTACGGAGT CTTCAGGCCC AGTTTCTCAC TTTAGCCAAT 4128 TCGAGGGCTC CTTGTGGTGG GATCAGAACT AATCCAGAGT GTGGGAAAGT 4178 GACAGTCAAA ACCCCACCTG GAGCAAATAA AAAAACATAC AAAACGTAAA 4278 AAAAAAAAA AAAAAA



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Tüle: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

### Human DNMT3B1 DNA Sequence:

1	GGCCGCGAAT	TCGGCACGAG	CCCTGCACGG	CCGCCAGCCG	GCCTCCCGCC
51	AGCCAGCCCC	GACCCGCGGC	TCCGCCGCCC	AGCCGCGCCC	CAGCCAGCCC
101	TGCGGCAGGA	AAGCATGAAG	GGAGACACCA	GGCATCTCAA	TGGAGAGGAG
151	GACGCCGGCG	GGAGGGAAGA	CTCGATCCTC	GTCAACGGGG	CCTGCAGCGA
201	CCAGTCCTCC	GACTCGCCCC	CAATCCTGGA	GGCTATCCGC	ACCCCGGAGA
251	TCAGAGGCCG	AAGATCAAGC	TCGCGACTCT	CCAAGAGGGA	GGTGTCCAGT
301	CTGCTAAGCT	ACACACAGGA	CTTGACAGGC	GATGGCGACG	GGGAAGATGG
351	GGATGGCTCT	GACACCCCAG	TCATGCCAAA	GCTCTTCCGG	GAAACCAGGA
401	CTCGTTCAGA	AAGCCCAGCT	GTCCGAACTC	GAAATAACAA	CAGTGTCTCC
451	AGCCGGGAGA	GGCACAGGCC	TTCCCCACGT	TCCACCCGAG	GCCGGCAGGG
501	CCGCAACCAT	GTGGACGAGT	CCCCCGTGGA	GTTCCCGGCT	ACCAGGTCCC
551	TGAGACGGCG	GGCAACAGCA	TCGCCAGGAA	CGCCATGGCC	GTCCCCTCCC
601	AGCTCTTACC	TTACCATCGA	CCTCACAGAC	GACACAGAGG	ACACACATGG
651	GACGCCCCAG	AGCAGCAGTA	CCCCCTACGC	CCGCCTAGCC	CAGGACAGCC
701	AGCAGGGGG	CATGGAGTCC	CCGCAGGTGG	AGGCAGACAG	TGGAGATGGA
751	GACAGTTCAG	AGTATCAGGA	TGGGAAGGAG	TTTGGAATAG	GGGACCTCGT
801	GTGGGGAAAG	ATCAAGGGCT	TCTCCTGGTG	GCCCGCCATG	GTGGTGTCTT
851	GGAAGGCCAC	CTCCAAGCGA	CAGGCTATGT	CTGGCATGCG	GTGGGTCCAG
901	TGGTTTGGCG	ATGGCAAGTT	CTCCGAGGTC	TCTGCAGACA	AACTGGTGGC
951	ACTGGGGCTG	TTCAGCCAGC	ACTITAATIT	GGCCACCTTC	AATAAGCTCG
001	TCTCCTATCG	AAAAGCCATG	TACCATGCTC	TGGAGAAAGC	TAGGGTGCGA
051	GCTGGCAAGA	CCTTCCCCAG	CAGCCCTGGA	GACTCATTGG	AGGACCAGCT
101	GAAGCCCATG	TTGGAGTGGG	CCCACGGGGG	CTTCAAGCCC	ACTGGGATCG
151	AGGGCCTCAA	ACCCAACAAC	ACGCAACCAG	TGGTTAATAÄ	GTCGAAGGTG

FIG. 1D-1

BY (C.6) 3 (C.4) (A.55)

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1201	CGTCGTGCAG	GCAGTAGGAA	ATTAGAATCA	AGGAAATACG	AGAACAAGAC
1251	TCGAAGACGC	ACAGCTGACG	ACTCAGCCAC	CTCTGACTAC	TGCCCGCAC
1301	CCAAGCGCCT	CAAGACAAAT	TGCTATAACA	ACGGCAAAGA	CCGAGGGGAT
1351	GAAGATCAGA	GCCGAGAACA	AATGGCTTCA	GATGTTGCCA	ACÁACAAGAG
1401	CAGCCTGGAA	GATGGCTGTT	TGTCTTGTGG	CAGGAAAAAC	CCCGTGTCCT
1451	TCCACCCTCT	CTTTGAGGGG	GGGCTCTGTC	AGACATGCCG	GGATCGCTTC
1501	CTTGAGCTGT	TTTACATGTA	TGATGACGAT	GGCTATCAGT	CTTACTGCAC
1551	TGTGTGCTGC	GAGGGCCGAG	AGCTGCTGCT	TTGCAGCAAC	ACGAGCTGCT
1601	GCCGGTGTTT	CTGTGTGGAG	TGCCTGGAGG	TGCTGGTGGG	CACAGGCACA
1651	GCGGCCGAGG	CCAAGCTTCA	GGAGCCCTGG	AGCTGCTACA	TGTGTCTCCC
1701	GCAGCGCTGT	CATGGCGTCC	TGCGGCGCCG	GAAGGACTGG	AACGTGCGCC
1751	TGCAGGCCTT	CTTCACCAGT	GACACGGGC	TTGAATACGA	AGCCCCCAAG
1801	CTGTACCCTG	CCATTCCCGC	AGCCCGAAGG	CGGCCCATTC	GAGTCCTGTC
1851	ATTGTTTGAT	GGCATCGCGA	CAGGCTACCT	AGTCCTCAAA	GAGTTGGGCA
1901	TAAAGGTAGG	AAAGTACGTC	GCTTCTGAAG	TGTGTGAGGA	GTCCATTGCT
1951	GTTGGAACCG	TGAAGCACGA	GGGGAATATC	AAATACGTGA	ACGACGTGAG
2001	GAACATCACA	AAGAAAAATA	TTGAAGAATG	GGGCCCATTT	GACTTGGTGA
2051	TTGGCGGAAG	CCCATGCAAC	GATCTCTCAA	ATGTGAATCC	AGCCAGGAAA
2101	GCCTGTATG	AGGGTACAGG	CCGCCTCTTC	TTCGAATTTT	ACCACCTGCT
2151	GAATTACTCA	CGCCCCAAGG	AGGGTGATGA	CCGGCCGTTC	TICTGGATGT
2201	TTGAGAATGT	TGTAGCCATG	AAGGTTGGCG	ACAAGAGGGA	CATCTCACGG
2251	TTCCTGGAGT	GTAATCCAGT	GATGATTGAT	GCCATCAAAG	TTTCTGCTGC
2301	TCACAGGGCC	CGATACTTCT	GGGGCAACCT	ACCCGGGATG	AACAGGCCCG
2351	TGATAGCATC	AAAGAATGAT	AAACTCGAGC	TGCAGGACTG	CTTGGAATAC
2401	AATAGGATAG	CCAAGTTAAA	GAAAGTACAG	ACAATAACCA	CCAAGTCGAA

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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

CTCGATCAAA CAGGGGAAAA ACCAACTTTT CCCTGTTGTC ATGAATGGCA 2451 AAGAAGATGT TTTGTGGTGC ACTGAGCTCG AAAGGATCTT TGGCTTTCCT 2501 GTGCACTACA CAGACGTGTC CAACATGGGC CGTGGTGCCC GCCAGAAGCT 2551 GCTGGGAAGG TCCTGGAGCG TGCCTGTCAT CCGACACCTC TTCGCCCCTC 2601 TGAAGGACTA CTTTGCATGT GAATAGTTCC AGCCAGGCCC CAAGCCCACT 2651 GGGGTGTGTG GCAGAGCCAG GACCCAGGAG GTGTGATTCC TGAAGGCATC 2701 CCCAGGCCCT GCTCTTCCTC AGCTGTGTGG GTCATACCGT GTACCTCAGT 2751 TCCCTCTTGC TCAGTGGGGG CAGAGCCACC TGACTCTTGC AGGGGTAGCC 2801 TGAGGTGCCG CCTCCTTGTG CACAAATCAG ACCTGGCTGC TTGGAGCAGC 2851 CTAACACGGT GCTCATTTTT TCTTCTCCTA AAACTTTAAA ACTTGAAGTA 2901 GGTAGCAACG TGGCTTTTTT TTTTTCCCTT CCTGGGTCTA CCACTCAGAG 2951 AAACAATGGC TAAGATACCA AAACCACAGT GCCGACAGCT CTCCAATACT 3001 CAGGITAATG CTGAAAAATC ATCCAAGACA GITATTGCAA GAGITTAATT 3051 TITGAAAACT GGGTACTGCT ATGTGTTTAC AGACGTGTGC AGTTGTAGGC 3101 3151 ATGTAGCTAC AGGACATTTT TAAGGGCCCA GGATCGTTTT TTCCCAGGGC AAGCAGAAGA GAAAATGTTG TATATGTCTT TTACCCGGCA CATTCCCCTT 3201 GCCTAAATAC AAGGGCTGGA GTCTGCACGG GACCTATTAG AGTATTTTCC 3251 3301 ACAATGATGA TGATTTCAGC AGGGATGACG TCATCATCAC ATTCAGGGCT 3351 ATTTTTCCC CCACAAACCC AAGGGCAGGG GCCACTCTTA GCTAAATCCC 3401 TCCCCGTGAC TGCAATAGAA CCCTCTGGGG AGCTCAGGAA GGGGTGTGCT 3451 GAGTICTATA ATATAAGCTG CCATATATIT TGTAGACAAG TATGGCTCCT 3501 CCATATCTCC CTCTTCCCTA GGAGAGGAGT GTGAAGCAAG GAGCTTAGAT AAGACACCCC CTCAAACCCA TTCCCTCTCC AGGAGACCTA CCCTCCACAG 3551 GCACAGGTCC CCAGATGAGA AGTCTGCTAC CCTCATTTCT CATCTTTTA 3601 3651 CTAAACTCAG AGGCAGTGAC AGCAGTCAGG GACAGACATA CATTTCTCAT

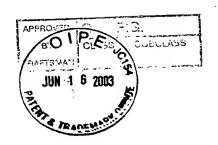
FIG. 1D-3



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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides

ACCTICCCCA CATCTGAGAG ATGACAGGGA AAACTGCAAA GCTCGGTGCT CCCTTTGGAG ATTTTTTAAT CCTTTTTTAT TCCATAAGAA GTCGTTTTTA 3751 GGGAGAACGG GAATTCAGAC AAGCTGCATT TCAGAAATGC TGTCATAATG 3801 GITITTAACA CCTTTTACTC ITCTTACTGG IGCTATITIG TAGAATAAGG 3851 AACAACGTTG ACAAGTTTTG TGGGGCTTTT TATACACTTT TTAAAATCTC 3901 AAACTICTAT TITTATGTIT AACGTTITCA TTAAAATTIT TITGTAACTG 3951 GAGCCACGAC GTAACAAATA TGGGGAAAAA ACTGTGCCTT GTTTCAACAG 4001 TITTIGCTAA TITTIAGGCT GAAAGATGAC GGATGCCTAG AGTITACCTT 4051 ATGTTTAATT AAAATCAGTA TITGTCTAAA AAAAAAAAAA AAAAA

FIG. 1D-4



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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides
and Uses Thereof

#### Mouse Dnmt3a Protein

MPSSGPGDTS SSSLEREDDR KEGEEQEENR GKEERQEPSA TARKVGRPGR KRKHPPVESS DTPKDPAVTT KSQPMAQDSG PSDLLPNGDL EKRSEPQPEE GSPAAGOKGG APAEGEGTET PPEASRAVEN GCCVTKEGRG ASAGEGKEQK 101 151 QTNIESMKME GSRGRLRGGL GWESSLRQRP MPRLTFQAGD PYYISKRKRD EWLARWKREA EKKAKVIAVM NAVEENQASG ESQKVEEASP PAVQQPTDPA 201 SPTVATTPEP VGGDAGDKNA TKAADDEPEY EDGRGFGIGE LVWGKLRGFS 251 WWPGRIVSWW MTGRSRAAEG TRWVMWFGDG KFSVVCVEKL MPLSSFCSAF 301 HQATYNKQPM YRKAIYEVLQ VASSRAGKLF PACHDSDESD SGKAVEVQNK 351 QMIEWALGGF QPSGPKGLEP PEEEKNPYKE VYTDMWVEPE AAAYAPPPPA 401 KKPRKSTTEK PKVKEIIDER TRERLVYEVR QKCRNIEDIC ISCGSLNVTL 451 501 EHPLFIGGMC QNCKNCFLEC AYQYDDDGYQ SYCTICCGGR EVLMCGNNNC CRCFCVECVD LLVGPGAAQA AIKEDPWNCY MCGHKGTYGL LRRREDWPSR 551 LOMFFANNHO QEFDPPKVYP PVPAEKRKPI RVLSLFDGIA TGLLVLKDLG 601 IQVDRYIASE VCEDSITVGM VRHQGKIMYV GDVRSVTQKH IQEWGPFDLV 651 IGGSPCNDLS IVNPARKGLY EGTGRLFFEF YRLLHDARPK EGDDRPFFWL 701 FENVVANGVS DKRDISRFLE SNPVMIDAKE VSAAHRARYF WGNLPGMNRP 751 LASTVNDKLE LQECLEHGRI AKFSKVRTIT TRSNSIKQGK DQHFPVFMNE 801 851 KEDILWCTEM ERVFGFPVHY TDVSNMSRLA RQRLLGRSWS VPVIRHLFAP 901 LKEYFACV\*

FIG. 2A

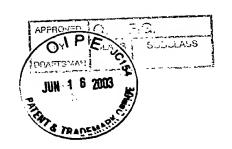


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and Uses Thereof

#### Mouse Dnmt3b1 Protein

MKGDSRHLNE EEGASGYEEC IIVNGNFSDQ SSDTKDAPSP PVLEAICTEP VCTPETRGRR SSSRLSKREV SSLLNYTQDM TGDGDRDDEV DDGNGSDILM PKLTRETKDT RTRSESPAVR TRHSNGTSSL ERQRASPRIT RGRQGRHHVQ 101 151 EYPVEFPATR SRRRRASSSA STPWSSPASV DFMEEVTPKS VSTPSVDLSQ DGDQEGMDTT QVDAESRDGD STEYQDDKEF GIGDLVWGKI KGFSWWPAMV 201 VSWKATSKRQ AMPGMRWVQW FGDGKFSEIS ADKLVALGLF SQHFNLATFN 251 KLVSYRKAMY HTLEKARVRA GKTFSSSPGE SLEDQLKPML EWAHGGFKPT 301 GIEGLKPNKK QPVVNKSKVR RSDSRNLEPR RRENKSRRRT TNDSAASESP 351 PPKRLKTNSY GGKDRGEDEE SRERMASEVT NNKGNLEDRC LSCGKKNPVS 401 451 FHPLFEGGLC OSCRDRFLEL FYMYDEDGYQ SYCTVCCEGR ELLLCSNTSC 501 CRCFCVECLE VLVGAGTAED AKLQEPWSCY MCLPQRCHGV LRRRKDWNMR 551 LODFFTTDPD LEEFEPPKLY PAIPAAKRRP IRVLSLFDGI ATGYLVLKEL GIKVEKYIAS EVCAESIAVG TVKHEGQIKY VNDVRKITKK NIEEWGPFDL 601 VIGGSPCNDL SNVNPARKGL YEGTGRLFFE FYHLLNYTRP KEGDNRPFFW 651 MFENYVAMKY NDKKDISRFL ACNPVMIDAI KVSAAHRARY FWGNLPGMNR 701 PVMASKNDKL ELQDCLEFSR TAKLKKVQTI TTKSNSIRQG KNQLFPVVMN 751 GKDDVLWCTE LERIFGFPAH YTDVSNMGRG ARQKLLGRSW SVPVIRHLFA 801 851 PLKDYFACE\*

FIG. 2B

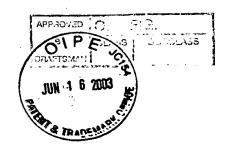


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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides
and Uses Thereof

#### Human DNMT3A Protein

MPAMPSSGPG DTSSSAAERE EDRKDGEEQE EPRGKEERQE PSTTARKVGR PGRKRKHPPV ESGDTPKDPA VISKSPSMAQ DSGASELLPN GDLEKRSEPQ 51 PFFGSPAGGO KGGAPAEGEG AAETLPEASR AVENGCCTPK EGRGAPAEAG 101 KFOKETNIFS MKMEGSRGRL RGGLGWESSL RORPMPRLTF QAGDPYYISK 151 RKRDEWLARW KREAEKKAKV IAGMNAVEEN QGPGESQKVE EASPPAVQQP 201 TDPASPTVAT TPEPVGSDAG DKNATKAGDD EPEYEDGRGF GIGELVWGKL 251 RGFSWWPGRI VSWWMTGRSR AAEGTRWVMW FGDGKFSVVC VEKLMPLSSF 301 CSAFHQATYN KQPMYRKAJY EVLQVASSRA GKLFPVCHDS DESDTAKAVE 351 401 VONKPMIEWA LGGFOPSGPK GLEPPEEEKN PYKEVYTDMW VEPEAAAYAP PPPAKKPRKS TAEKPKVKEI IDERTRERLV YEVRQKCRNI EDICISCGSL 451 NVTLEHPLFV GGMCQNCKNC FLECAYQYDD DGYQSYCTIC CGGREVLMCG 501 NNNCCRCFCV ECVDLLVGPG AAQAAIKEDP WNCYMCGHKG TYGLLRRRED 551 WPSRLQMFFA NNHDQEFDPP KVYPPVPAEK RKPIRVLSLF DGIATGLLVL 601 KDLGIQVDRY [ASEVCEDS] TVGMVRHQGK IMYVGDVRSV TQKHIQEWGP 651 FDLVIGGSPC NDLSIVNPAR KGLYEGTGRL FFEFYRLLHD ARPKEGDDRP 701 FFWLFENVVA MGVSDKRDIS RFLESNPVMI DAKEVSAAHR ARYFWGNLPG 751 MNRPLASTVN DKLELQECLE HGRIAKFSKV RTITTRSNSI KQGKDQHFPV 801 FMNEKEDILW CTEMERVFGF PVHYTDVSNM SRLARQRLLG RSWSVPVIRH 851 901 LFAPLKEYFA CV\*

FIG. 2C

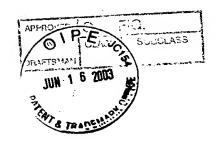


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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides
and Uses Thereof

#### Human DNMT3B1 Protein

MKGDTRHLNG EEDAGGREDS ILVNGACSDQ SSDSPPILEA IRTPEIRGRR SSSRLSKREV SSLLSYTODL TGDGDGEDGD GSDTPVMPKL FRETRTRSES PAVRTRNINS VSSRERHRPS PRSTRGROGR NHVDESPVEF PATRSLRRRA TASAGTPWPS PPSSYLTIDL TDDTEDTHGT PQSSSTPYAR LAQDSQQGGM 151 ESPOVEADSG DGDSSEYQDG KEFGIGDLVW GKIKGFSWWP AMVVSWKATS 201 251 KROAMSCMRW VOWFGDGKFS EVSADKLVAL GLFSQHFNLA TFNKLVSYRK AMYHALEKAR VRAGKTFPSS PGDSLEDQLK PMLEWAHGGF KPTGIEGLKP 301 NNTOPYVNKS KVRRAGSRKL ESRKYENKTR RRTADDSATS DYCPAPKRLK 351 TNCYNNGKDR GDEDQSREQM ASDVANNKSS LEDGCLSCGR KNPVSFHPLF 401 EGGLCOTCRD RFLELFYMYD DDGYQSYCTV CCEGRELLLC SNTSCCRCFC 451 VECLEVLYGT GTAAEAKLQE PWSCYMCLPQ RCHGVLRRRK DWNVRLQAFF 501 TSDTGLEYEA PKLYPAIPAA RRRPIRVLSL FDGIATGYLV LKELGIKVGK 551 YVASEVCEES IAVGTVKHEG NIKYVNDVRN ITKKNIEEWG PFDLVIGGSP 601 CNDLSNVNPA RKGLYEGTGR LFFEFYHLLN YSRPKEGDDR PFFWMFENVV 651 AMKVGDKRDI SRFLECNPVM IDAIKVSAAH RARYFWGNLP GMNRPVIASK 701 NDKLELQDCL EYNRIAKLKK VQTITTKSNS IKQGKNQLFP VVMNGKEDVL 751 WCTELERIFG FPVHYTDVSN MGRGARQKLL GRSWSVPVIR HLFAPLKDYF 801 ACE\* 851

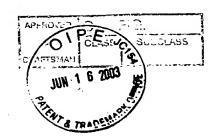
FIG. 2D



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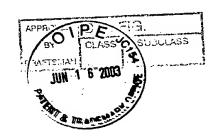
Dnmt3a	1	MPSSGPGDTSSSSLEREDDRKEGEEQEENRGKEERQEPSATARKVGRPGR	50
Dnmt3a		KRKHPPVESSDTPKDPAVTTKSQPMAQDSGPSDLLPNGDLEKRSEP	
Dnmt3b	1	.   . : :     . :      MKGDSRHLNEEEGASGYEECIIVNGNFSDQSSD	33
Dnmt3a	97	QPEEGSPAAGQKGGAPAEGEGTETPPEAS.RAVENGCCVTKEGR	139
Dnmt3b	34	:      .     .       .   .   TKDAPSPPVLEAICTEPVCTPETRGRRSSSRLSKREVSSLLNYTQDMTGD	83
Dnmt3a	140	GASAGEGKEQKQTNIESMKMEGSRGRLRGGLGWESSLRQ	178
Dnmt3b	84		133
Dnmt3a	179	RPMPRLTFQAGDPYYISKRKRDEWLARWKREAEKKAKVIAVMNAVEENQA	228
Dnmt3b	134	:  :::  : :: :   RASPRITRGRQGRHHVQEYPVEFPATRSRRRASSSASTPWSSPA	178
Dnmt3a	229	SGESQKVEEASPPAVQQPTDPASPTVATTPEPVGGDAGDKNATKAADDEP	278
Dnmt3b	179	: .  .  .   :     .    SVDFMEEVTPKSVSTPSVDLSQDGDQEGMDTTQVDAESRDGDST	222
Dnmt3a	279	EYEDGRGFGIGELVWGKLRGFSWWPGRIVSWWMTGRSRAAEGTRWVMWFG	328
Dnmt3b	223	EYQDDKEFGIGDLVWGKIKGFSWWPAMVVSWKATSKRQAMPGMRWVQWFG	272
Dnmt3a	329	DGKFSVVCVEKLMPLSSFCSAFHQATYNKQPMYRKAIYEVLQVASSRAGK	378
Dnmt3b	273	DGKFSEISADKLVALGLFSQHFNLATFNKLVSYRKAMYHTLEKARVRAGK	322
Dnmt3a	379	LFPACHDSDESDSGKAVEVQNKQMIEWALGGFQPSGPKGLEPPEEEKN	426
Dnmt3b		TFSSSPGESLEDQLKPMLEWAHGGFKPTGIEGLKPNKKQPVVN	
Dnmt3a	427	PYKEVYTDMW.VEPEAAAYAPPPPAKKPRKSTTEKPK	462
Dnmt3b	366	KSKVRRSDSRNLEPRRRENKSRRRTTNDSAASESPPPKRLKTNSYGGKDR	415

FIG.3A-1



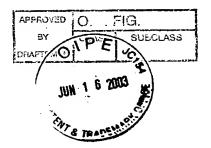
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Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

Dnmt3a	463	VKEIIDERTRERLVYEVRQKCRNIEDICISCGSLNVTLEHPFFIGGMCQN .    .  :     :              :   .	512
Dnmt3b	416	GEDEESRERMASEVTNNKGNLEDRCLSCGKKNPVSFHPLFEGGLCQS	462
Dnmt3a	513	CKNCFLECAYQYDDDGYQSYCTICCGGREVLMCGNNNCCRCFCVECVDLL	562
Dnmt3b	463	CRDRFLELFYMYDEDGYQSYCTVCCEGRELLLCSNTSCCRCFCVECLEVL	512
Dnmt3a	563	VGPGAAQAAIKEDPWNCYMCGHKGTYGLLRRREDWPSRLQMFFANNHD.Q	611
Dnmt3b	513	VGAGTAEDAKLQEPWSCYMCLPQRCHGVLRRRKDWNMRLQDFFTTDPDLE	562
Dnmt3a	612	EFDPPKVYPPVPAEKRKPIRVLSLFDGIATGLLVLKDLGIQVDRYIASEV	661
Dnmt3b	563	EFEPPKLYPAIPAAKRRPIRVLSLFDGIATGYLVLKELGIKVEKYIASEV	612
Dnmt3a	662	CEDSITVGMVRHQGKIMYVGDVRSVTQKHIQEWGPFDLVIGGSPCNDLSI	711
Dnmt3b	613	CAESIAVGTVKHEĞQİKYVNDVRKITKKNİEEWGPFDLVİĞĞSPCNDLSN	662
Dnmt3a	712	VNPARKGLYEGTGRLFFEFYRLLHDARPKEGDDRPFFWLFENVVAMGVSD	761
Dnmt3b	663	VNPARKGLYEGTGRLFFEFYHLLNYTRPKEGDNRPFFWMFENVVAMKVND	712
Dnmt3a		KRDISRFLESNPVMIDAKEVSAAHRARYFWGNLPGMNRPLASTVNDKLEL	
Dnmt3b		KKDISRFLACNPVMIDAIKVSAAHRARYFWGNLPGMNRPVMASKNDKLEL	
Dnmt3a		QECLEHGRIAKFSKVRTITTRSNSIKQGKDQHFPVFMNEKEDILWCTEME	
Dnmt3b		QDCLEFSRTAKLKKVQTITTKSNSIRQGKNQLFPVVMNGKDDVLWCTELE	
Dnmt3a		RVFGFPVHYTDVSNMSRLARQRLLGRSWSVPVIRHLFAPLKEYFACV*	909
Dnmt3b	813	RIFGFPAHYTDVSNMGRGARQKLLGRSWSVPVIRHLFAPLKDYFACE*	860



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Title: De/Novo DNA Cytosine Methyltransferase Genes, Polypeptides
and Uses Thereof

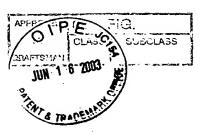
DNMT3A	1 MPAMPSSGPGDTSSSAAEREEDRKDGEEQEEPRGKEERQEPSTTARKVGR
DNMT3A	51 PGRKRKHPPVESGDTPKDPAVISKSPSMAQDSGASELLPNGDLEKRSEPQ
DNMT3B	: :  : 1MKGDTRHLNGEEDAGGREDSILVNGACSDQSSDSP
DNMT3A	101 PEEGSPAGGQKGGAPAEGEGAAETLPEASRAVENGCCTPKEGRGAPAEAG
DNMT3B	. :     . : :   .: 36 PILEAIRTPEIRGGWASSRLSKREVSSLLSYTQDLTGDGDGEDGDGSDTP
DNMT3A	151 KEQKETNIESMKMEGSRGRLRGGLGWESSLRQRPMPRLTFQAGDPYYISK
DNMT3B	86 VMPKLFRETRTRSESPAVRTRNNNSVSSRERHRPSPRSTRGRQGRNHVDE
DNMT3A	201 RKRDEWLARWKREAEKKAKVIAGMNAVEENQGPGESQKVEEASPPAVQQP
DNMT3B	136 SPVEFPATRSLRRRATASAGTPWPSPPSSYLTIDLTDDTEDTHGTPQS
DNMT3A	251 TDPASPTVATTPEPVGSDAGDKNATKAGDDEPEYEDGRGFGIGELVWGKL
DNMT3B	184 SSTPYARLAQDSQQGGMESPQVEADSGDGDSSEYQDGKEFGIGDLVWGKI
DNMT3A	301 RGFSWWPGRIVSWWMTGRSRAAEGTRWVMWFGDGKFSVVCVEKLMPLSSF:
DNMT3B	234 KGFSWWPAMVVSWKATSKRQAMSGMRWVQWFGDGKFSEVSADKLVALGLF
DNMT3A	351 CSAFHQATYNKQPMYRKAIYEVLQVASSRAGKLFPVCHDSDESDTAKAVE
DNMT3B	
DNMT3A	401 VQNKPMIEWALGGFQPSGPKGLEPPEEEKNPYKEVYTDMWVE
DNMT3B	327 DQLKPMLEWAHGGFKPTGIEGLKPNNTQPVVNKSKVRRAGSRKLESRKYE
DNMT3A	443PEAAAYAPPPPAKKPRKSTAEKPKVKEIIDERTRERLVYEVRQ
DNMT3B	377 NKTRRTADDSATSDYCPAPKRLKTNCYNNGKDRGDEDQSREQMASDVAN



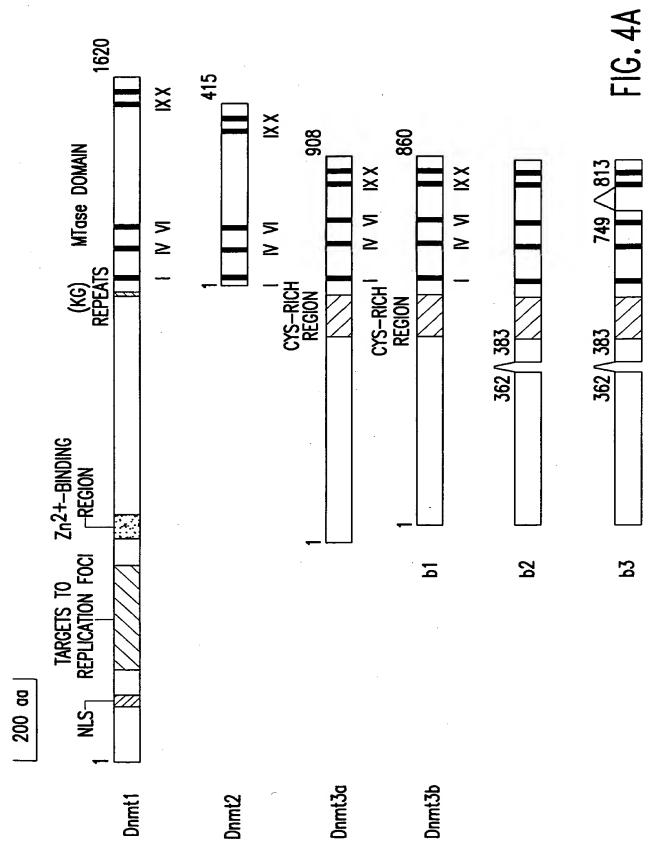
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Inventors: Li et al.; Tel: 202/371-2600
Täle: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

DNMT3A	486	KCRNIEDICISCGSLNVILEHPLFVGGMCQNCKNCFLECAYQYDDDGYQS
DNMT3B	427	NKSSLEDGCLSCGRKNPVSFHPLFEGGLCQTCRDRFLELFYMYDDDGYQS
DNMT3A	536	YCTICCGGREVLMCGNNNCCRCFCVECVDLLVGPGAAQAAIKEDPWNCYM
DNMT3B	477	:      . :    .        .:.        ::  .    YCTVCCEGRELLLCSNTSCCRCFCVECLEVLVGTGTAAEAKLQEPWSCYM
DNMT3A	586	CGHKGTYGLLRRREDWPSRLQMFFANNHDQEFDPPKVYPPVPAEKRKPIR
DNMT3B	527	. : .             ::   .   : :    CLPQRCHGVLRRRKDWNVRLQAFFTSDTGLEYEAPKLYPAIPAARRRPIR
DNMT3A	636	VLSLFDGIATGLLVLKDLGIQVDRYIASEVCEDSITVGMVRHQGKIMYVG
DNMT3B	577	
DNMT3A	686	DVRSVTQKHIQEWGPFDLVIGGSPCNDLSIVNPARKGLYEGTGRLFFEFY
DNMT3B	627	.: . :
DNMT3A	736	RLLHDARPKEGDDRPFFWLFENVVAMGVSDKRDISRFLESNPVMIDAKEV
DNMT3B	677	
DNMT3A	786	SAAHRARYFWGNLPGMNRPLASTVNDKLELQECLEHGRIAKFSKVRTITT
DNMT3B	727	SAAHRARYFWGNLPGMNRPVIASKNDKLELQDCLEYNRIAKLKKVQTITT
DNMT3A	836	RSNSIKQGKDQHFPVFMNEKEDILWCTEMERVFGFPVHYTDVSNMSRLAR:
DNMT3B	777	KSNSIKQGKNQLFPVVMNGKEDVLWCTELERIFGFPVHYTDVSNMGRGAR
DNMT3A	886	QRLLGRSWSVPVIRHLFAPLKEYFACV*
DNMT3B	827	:

FIG.3B-2



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APPROVED O. FIG.
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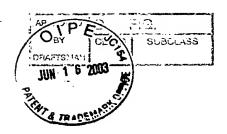
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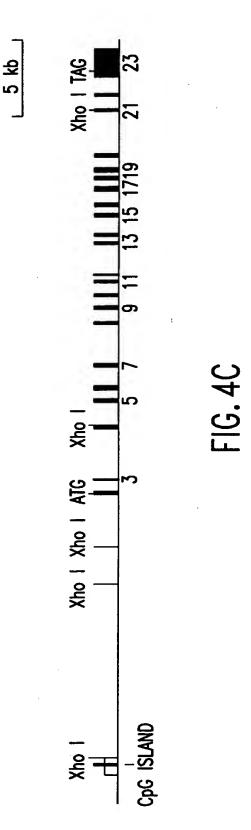
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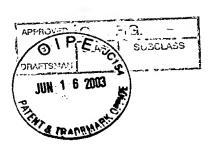
FIG. 4B

DNMT3A DNMT3B Zmt3



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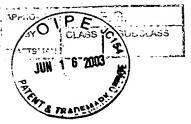
Appl. No. 09/720,086; 102(e): July 23, 2001
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Inventors: Li et al.; Tel: 202/371-2600
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and Uses Thereof

.cattttgttctccagTTAAAG . CTGTTTCCTCTACAGGCCGAA .tgtttccttataaagGACTTG .tecttetgtecacagTCCCTG .ctctctggctgccagGATCGC .ctctggcccccacagGCTACC .cttttctgagcacagAGGGTA .tggcttctcccacagGAAAGC .ctctcttgcttctagGTCCGA .tgggttttcttccagGATGGG .gtctttctctttagGTCTCT .ttttcccctcaaaagTGGTTA .aattacctttcacagAGAACA .ttcttttctcaatagAACAAA .ttccttacctggcagGAAGCC .gtctctctttcagATTGAA .tctttctccccacaqTGTAAT .tttggctgttcccagGCCCGT .ctccggtacccccagGATCTT .actctgcctttgcagAAAGCT .ctgtttttcttacagATGGCT .tgccactgggtccagGTGTTT 801bp) 602bp) 917bp) 765bp) 1813bp) 115bp) 417bp) (dq009 824bp) 536bp) 352bp) 958bp) 887bp) 1160bp) (>=90bp) CGGCAGgtgagcgcccgggg.intron(17618bp) 1642bp) 2867bp) 1434bp) 1403bp) 2588bp) 1095bp) 3343bp) TATCAGgtatggccgagaggg.intron 60bp) AATACGgtatttccttcctgt.intron GCCGAGgtgattgttgggtac.intron TGGAAGgtaacgttctctccc.intron TGCCGGgtaagtcctcctact.intron CTGCCGgtgagcactgggccc.intron 184bp) GAATACgtaagccacaggctc.intron CGACAGgtgagttcgggggaac.intron 146bp) AAAAATgtgaggggagtctgt.intron TGTATGgtgagcatccttctc.intron CTGGAGgtgagggaatctggg.intron GAACAGgtaacaaagggctct.intron 70bp) GCCAAGttaaagaaagtacag.intron CGAAAGgtgagcaaggctgca.intron TCAGAGgtggctgggcagtgg.intron 62bp) ACACAGgtatggtctctgctc.intron CCAGCTgtaagtagccacacc.intron 125bp) ACCAGGgttgttccccagatg.intron TCCGAGgtgagtccggggaag.intron CTGGAGgtaacatgggatgag.intron 145bp) AACCAGgtgggaatgagtccc.intron [29bp] 126bp)( 45bp) r 80bp), 85bp) ( 91bp)' 222bp) 108bp) 113bp) 102bp) 149bp) 86bp) 148bp) 119bp) Exon23 (1585bp Exon20 3xon16 Exon19 Exon10 Exon11 Exon12 Exon13 Exon14 Exon15 Exon17 Exon18 3xon21 Exon22 Exon7 Exon9 Exon8 Exon5 Exon6 Exon2 Exon4 3xon1 Exon3

## FIG.4D

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## FIG. 5A

DRPFFWLFENVVAMGVSDK DRPFFWMFENVVAMKVGDK NRPFFWMFENVVAMKVNDK PQPFFWLFENVTFMQTHVK	PFDLVIGGSPCNDLSIVNP PFDLVIGGSPCNDLSNVNP PFDLLIGGSPCNDLSIVNP PFDLLIGGSPCNDLSIVNP	SLFDGIATGLLVLKDLG SLFDGIATGYLVLKELG SLFDGIATGYLVLKELG SLFDGIATGYLVLRDLG F-GG	Onmt3a DNMT3B Dnmt3b Zmt3 consensus
DRPFFWLFENVVAMGVSDK	PFDLVIGGSPONDLSIVNP	SLEDGIATGLLVLKDLG	Dnmt3a DNMT2B
DRPFFWLFENWVAMGVSDK	PFDLVIGGSPCNDLSIVNP	SLFDGIATGLLVLKDLG	DNMT3A
KOPKFFVFENVKGL INHDK	<b>EFDLLVGGSPCQSFSVAGH</b>	SLFSGIGAFEAALRNIG	M.Spr
KLPKYILLENWKGFEVSST	SFNMILMSPPCOPFIRIGL	ELYSGIGGMHHALRESH	Drumt2
YKPRFVLLENWKGLITTKL	EVDFIYGGPPCQGFSGVNR	DIFAGCGGLTLGLDLSG	Masc2
VRPRLFTVE ETDG IMDRQS	HVDILHLSPPCOTFSRAHT	<b>OTFCGGGGVSLGARQAG</b>	Masc1
FRPRYFLLENWRTFVSFNK	QVDFINGGPPCQGFSGMINR	DIFAGCGGLSHGLKKAG	MET1(Ath)
YRPRFFLLENWRNFVSYRR	<b>DVEMLCGGPPCQGFSGMNR</b>	<b>DVFSGCGGLSEGFHQAG</b>	Dnmt1
YRPRFFLLENVRNFVSFKR	<b>DVEMLCGGPPCQGFSGMNR</b>	DVFSGCGGLSEGFHQAG	DNMT1

GGFP FFGNILDRHRQVGNAVFPPL GGFP FAGNINHKHRQIGNAVPPPL GGFP FVGTLTDKRIIGNAVPPPL GGFP GLGGVKKWHRNIGNAVPVPL GGFP EKTTVKQRYRLLGNSLNVHV QAFD AGISNSQLYKQTGNSITVTV FGFP SNMSRLARQRLLGRSWSVPV FGFP SNMSRLARQRLLGRSWSVPV

DNMT1
Drmt1
Masc1
Masc2
Drmt2
M.Spr
DNMT3A
Dnmt3a
DNMT3B

RRLIPLECHRUGAFD AGIS
DILWCTEMERVFGFP SNMS
DILWCTELERIFGFP SNMS
DVLWCTELERIFGFP SNMC
DVLWCTELERIFGFP SNMC
DHIWITELERIFGFP KSMC

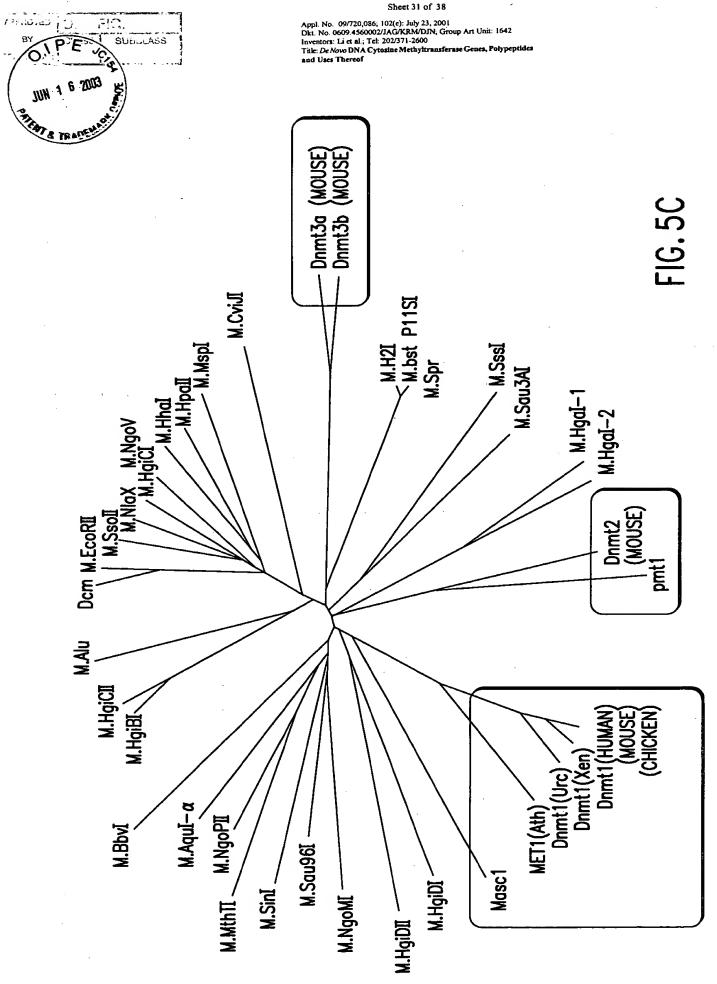
mt3 onsensus

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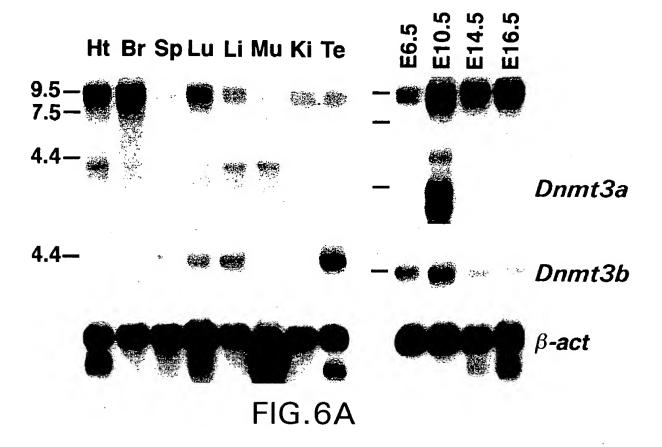
Appl. No. 09/720,086; 102(e): July 23, 2001
Dkt. No. 0609.4560002/JAG/KRM/DJN; Group Art Unit: 1642
Inventors: Li et al.; Tel: 202/371-2600
Title: De Novo DNA Cytosine Methyltransferase Genes, Polypeptides and Uses Thereof

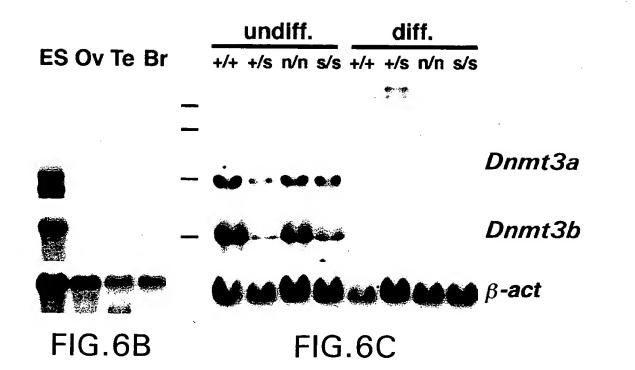
# FIG.5B

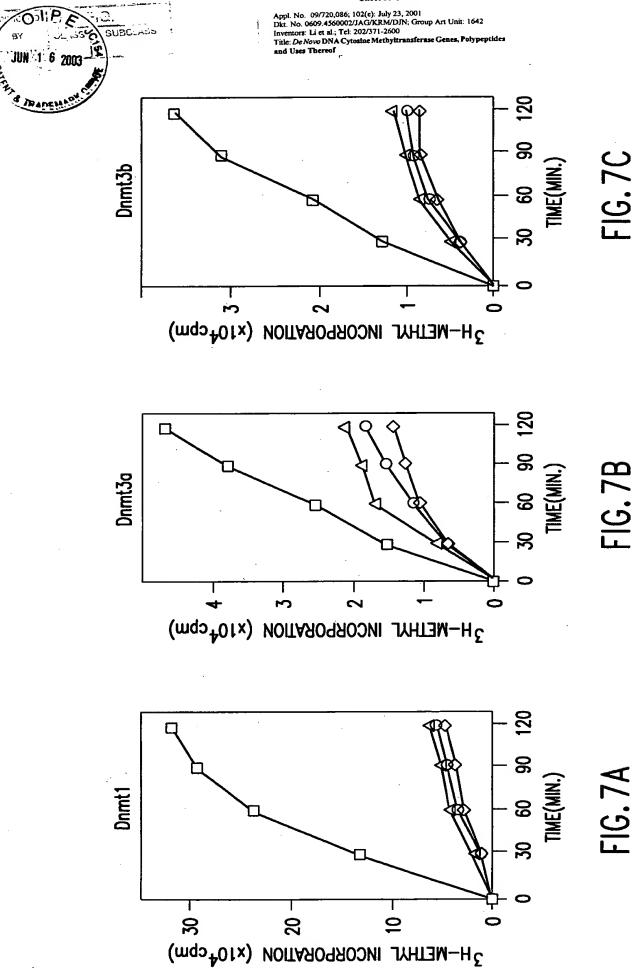




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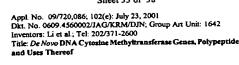


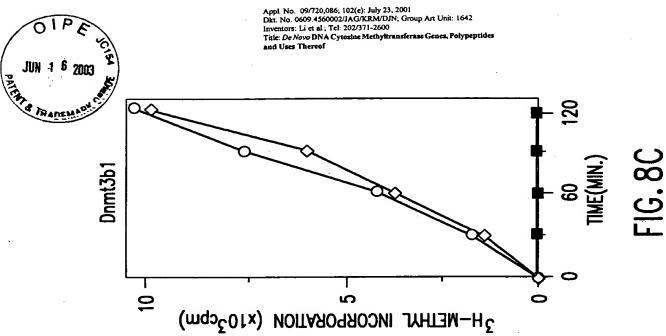
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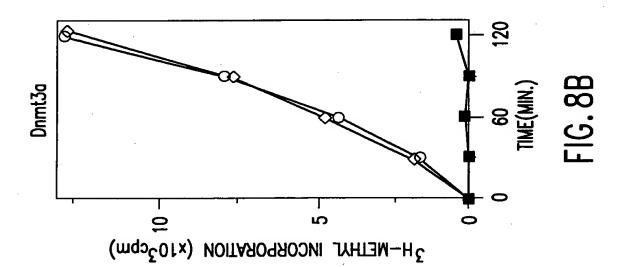
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67	5th	K/W	
<b>Dnmt3b</b>	¥	1st 3rd 5th	
Dnr	5th	K/W	
<b>Dnmt3a</b>	K H	tet 3rd 5th	
Dul	5th	K/W	
_	₹	tet 3rd ftb	
Dnmt	5th	K\W K\H K\H K\W	
<b>PMMLV</b>		K\H K\W K	
			1.0

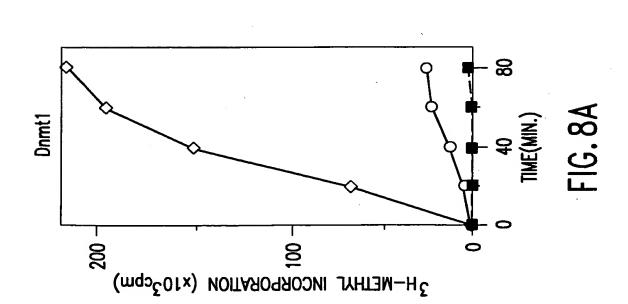
FIG.7D

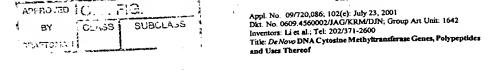


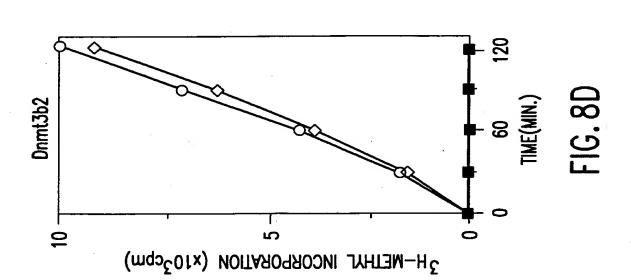


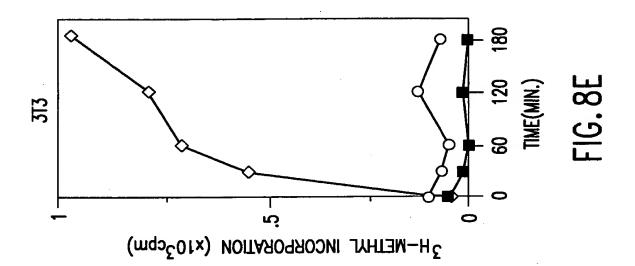




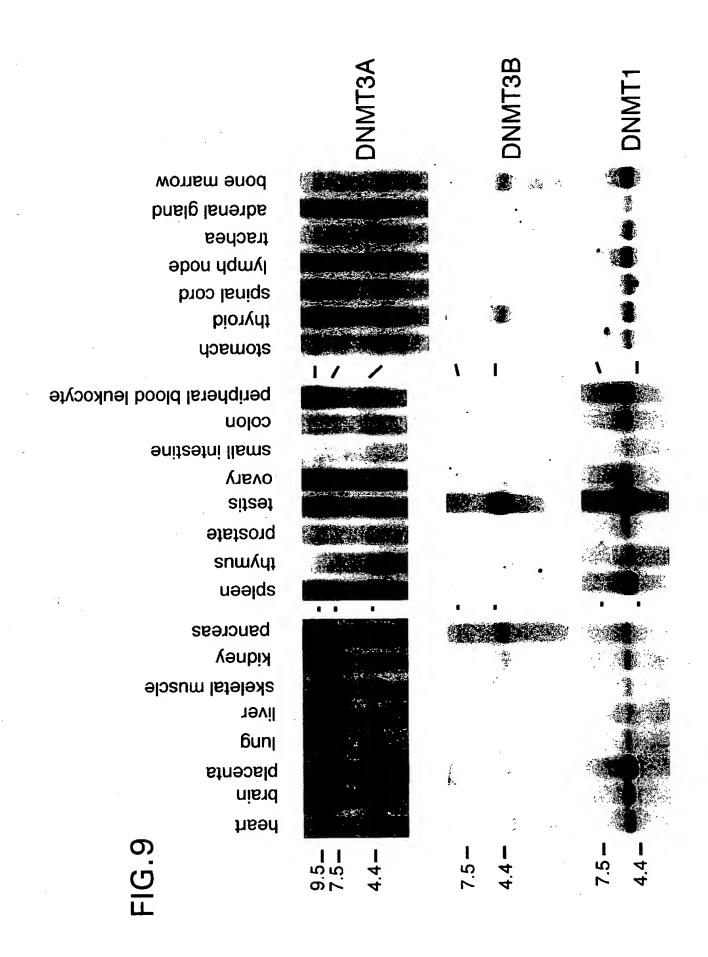








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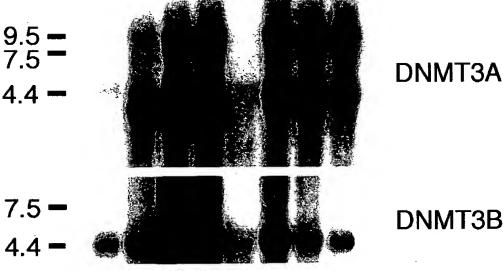


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promyelocytic leukemia HL-60

chronic myelogenous leukemia K-562 colorectal adenocarcinoma SW480 lymphoblastic leukemia MOLT-4 Burkitt's lymphoma Raji lung carcinoma A549 Hela cell S3

melanoma G361



DNMT3B

7.5 **-DNMT1** 4.4



b-Actin